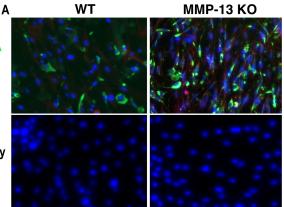
## **Supplementary Material**

## The Role of Matrix Metalloproteinase-13 (MMP13) in TGFβ/BMP Pathway Regulation of Fibro-Adipogenic Progenitor (FAP) Differentiation

Mengyao Liu<sup>a,b,c</sup> Brian T. Feeley<sup>a,b</sup> Hubert T. Kim<sup>a,b</sup> Xuhui Liu<sup>a,b</sup>

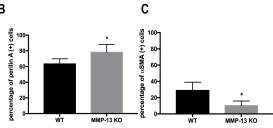
<sup>a</sup>San Francisco Veterans Affairs Health Care System, Department of Veterans Affairs, San Francisco, CA, USA, <sup>b</sup>Department of Orthopaedic Surgery, University of California, San Francisco, CA, USA, <sup>c</sup>California Northstate University, College of Medicine, Elk Grove, CA, USA **Perilipin A** aSMA

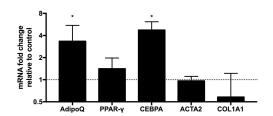
No primary Antibody Control



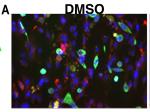


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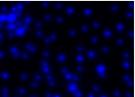


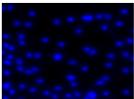


Supplemental Figure 1. FAPs from MMP13 KO mice have increased spontaneous adipogenesis and decreased spontaneous fibrogenesis compared to FAPs from WT mice cultured in the adipogenic medium. A) A typical image of immunostaining for perilipin A and  $\alpha$ SMA in FAPs from WT and MMP13 KO mice after 2 weeks of culturing in adipogenic medium. B) FAPs from MMP13 KO mice had significantly higher percentage of perilipin A positive cells compared to FAPs from WT mice. C) FAPs from MMP13 KO mice had a significantly lower percentage of  $\alpha$ SMA positive cells compared to FAPs from WT mice. D) Real time PCR showed that FAPs from MMP13 KO mice had a significantly higher expression of Adiponectin and C/EBP compared to FAPs from wildtype mice (\* p<0.05). Perilipin A aSMA

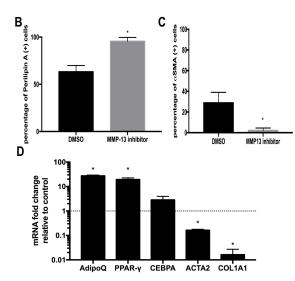


## MMP-13 Inhibitor





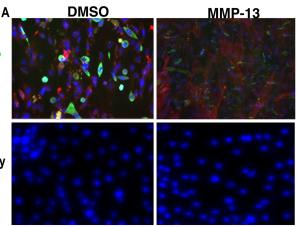




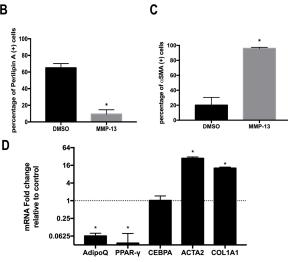
Supplemental Figure 2. Wildtype FAP treated with the MMP13 inhibitor had significantly increased adipogenesis in adipogenic medium. A) A typical image of immunostaining in FAPs treated with 10  $\mu$ M MMP13 inhibitor and 0.1% DMSO in the adipogenic medium for 2 weeks. B) WT FAPs treated with the MMP13 inhibitor had a significantly higher percentage of perilipin A (+) cells compared to those treated with DMSO. C) FAPs treated with the MMP13 inhibitor had a significantly reduced the number of  $\alpha$ SMA positive cells compared to DMSO. D) Real time PCR showed that FAPs treated with the MMP13 inhibitor had a significantly increased expression of Adiponectin, PPAR $\gamma$ , and decreased expression of  $\alpha$ SMA and Collagen I (\* p<0.05).

**Perilipin A** aSMA

No primary Antibody Control



В



Supplemental Figure 3. MMP13 treatment inhibits FAP adipogenesis and promotes FAP fibrogenesis in adipogenic medium. A) A typical image of immunostaining for FAPs treated with 100ng/ml MMP13 and 0.1% DMSO in adipogenic medium for 2 weeks. B) FAPs treated with MMP13 have a significantly reduced the number of perilipin (+) cells compared to DMSO. C) FAPs treated with MMP13 have a significantly increased the number of  $\alpha$ SMA (+) cells compared to the DMSO treatment group. D) Real time PCR results of FAPs treated with MMP13 showed a significantly decreased expression of adipogenesis-related genes (\* p<0.05).